## Amendments to the Claims

The listing of claims will replace all prior versions, and listings of claims in the application.

- 1. (currently amended) A method for isolation of biological macromolecules, said method comprising contacting a filtration apparatus multi-layer filter with a biological sample comprising said biological macromolecules, said filtration apparatus multi-layer filter comprising a first filter layer on top of a second filter layer such that said first filter layer is contacted with said biological macromolecules before said second filter layer, and said first filter layer having a pore size smaller than said second filter layer.
- 2. (original) The method of claim 1, wherein said biological sample is a cellular lysate.
- 3. (original) The method of claim 2, wherein said cellular lysate is derived from eukaryotic cells.
- 4. (original) The method of claim 2, wherein said cellular lysate is derived from prokaryotic cells.
- 5. (original) The method of claim 3, wherein said eukaryotic cells are selected from the group consisting of fungi, fish cells, yeast cells, plant cells and animal cells.

	6. (original)	The method of claim 1, wherein said biological macromolecules	
are nucleic acid molecules.			
are pro	7. (original) stein molecules	The method of claim 1, wherein said biological macromolecules	
RNA r	8. (original) molecules.	The method of claim 6, wherein said nucleic acid molecules are	
moleci		The method of claim 8, wherein said RNA molecules are mRNA	
DNA 1	10. (original) molecules.	The method of claim 6, wherein said nucleic acid molecules are	
or plas		The method of claim 10, wherein said DNA molecules are vectors	
	13. (cancelled)		
	14. (cancelled	)	

15. (cancelled)

- 16. (currently amended) The method of claim 1, wherein said pore size of said second filter <u>layer</u> is about 1  $\mu m$  to 500  $\mu m$ .
- 17. (currently amended) The method of claim 16, wherein said pore size of said second filter <u>layer</u> is about 10  $\mu$ m to 70  $\mu$ m.
- 18. (currently amended) The method of claim 17, wherein said pore size of said second filter <u>layer</u> is about 20 μm.
  - 19. (cancelled)
  - 20. (cancelled)
- 21. (currently amended) The method of claim 1, wherein said first filter <a href="layer">layer</a> comprises pores of sufficient size to retard the flow of cellular debris and particles.
- 22. (currently amended) The method of claim 21, wherein said pores of said first filter <u>layer</u> are about  $0.1~\mu m$  to  $1.0~\mu m$  in diameter.
- 23. (currently amended) The method of claim 21, wherein said pores of said first filter <u>layer</u> are about  $0.2 \mu m$  in diameter.

- 24. (currently amended) The method of claim 1, wherein said second filter <u>layer</u> is comprised of glass fibers, silica, paper, cellulose, nitrocellulose, diatomaceous earth, and acetylated cellulose.
- 25. (currently amended) The method of claim 1, wherein said first filter layer is comprised of one or more materials selected from the group consisting of hydrophobic polysolfone, hydrophilic polyether sulfone, cellulose, acetylated cellulose, nitrocellulose, polyester, polyolefin, scintered polyethylene, porous ceramics, silica, polypropylene, paper, and polysaccharide.
  - 26. (cancelled)
- 27. (currently amended) The method of claim 26, wherein said first filter layer has an average pore size of about 0.2  $\mu$ m, and said second filter layer has an average pore size of about 20  $\mu$ m.
- 28. (currently amended) The method of claim 1, wherein said first filter layer is provided in a form selected from the group consisting of wafer, cylindrical, rectangular, beads, gels, square, cartridge, swab tip, plug, frit, membrane, sheets or inserts.
- 29. (currently amended) The method of claim 1, wherein said filtration apparatus multi-layer filter is provided in a form that is suitable to be inserted into a tube, microspin tube, microfuge tube, spin cartridge, vial, ampule, bag or suitable to fit multi-well plates typically used in processing of multiple samples, including, 6-well plates, 12-

well plates, 24-well plates, 48-well plates, 96-well plates, 384-well plates, and the like, or suitable to fit into other plate sizes such as 35 mm plates, 60 mm plates, 100 mm plates, or 150 mm plates, and the like.

- 30. (original) The method of claim 1, wherein the flow of the sample is facilitated by centrifugation, gravity, pressure, vacuum, or any combination thereof.
- 31. (currently amended) A method for isolation of biological macromolecules, said method comprising;
- (a) contacting cells or cellular source containing the macromolecules of interest with a composition capable of lysing all or substantially all of said cells to give a lysate; and
- (b) contacting the lysate with a filtration apparatus multi-layer filter, wherein the apparatus comprises two filters layers, with a first filter layer on top of a second filter layer such that said first filter layer is contacted with said lysate before said second filter layer, and said first filter layer having a pore size smaller than said second filter layer; and
- (c) promoting the flow through the filtration apparatus multi-layer filter.

## 32. -54. (cancelled)

55. (currently amended) A process for isolating biological macromolecules comprising, separating a lysed natural source in a sample by filtration, wherein said

sample is passed through a <u>filtration apparatus</u> <u>multi-layer filter</u> comprising a first filter <u>layer</u> on top of a second filter <u>layer</u> such that said first filter <u>layer</u> is contacted with said biological macromolecules before said second filter <u>layer</u>, and said first filter <u>layer</u> having a pore size smaller than said second filter <u>layer</u>.

56. (currently amended) The process according to claim 55, wherein the flow through the <u>filtration apparatus multi-layer filter</u> is promoted by applying positive or negative pressure, or by gravity, or by gravity increased by centrifugation, or by a combination thereof.

57. (currently amended) The process according to claim 55, wherein said biological macromolecule is a plasmid DNA or genomic DNA having a size of from 1 to 50 kb.

- 58. (cancelled)
- 59. (cancelled)
- 60. (cancelled)
- 61. (currently amended) The process according to claim 55, wherein said first filter <u>layer</u> has a pore size of 0.1 to 1.0  $\mu$ m, and the second filter <u>layer</u> has a pore size of 1 to 500  $\mu$ m.
- 62. (currently amended) The process according to claim 55, wherein said first filter <u>layer</u> comprise one or more materials selected from the group consisting of hydrophobic polysolfone, hydrophilic polyether sulfone, cellulose, acetylated cellulose,

nitrocellulose, polyester, polyolefin, scintered polyethylene, porous ceramics, silica, polypropylene, paper, and polysaccharide.

63. (currently amended) The process according to claim 55, wherein said second filter <u>layer</u> is comprised of glass fibers, silica, paper, cellulose, nitrocellulose, diatomaceous earth, and acetylated cellulose.

64-65. (cancelled)

66. (currently amended) The method of any of claims 1, 31, or 55, wherein said second filter <u>layer</u> shears genomic DNA.